

-continued

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1.-38. (canceled)

39. A method for detecting mutations from exposure to a substance, the method comprising:

- (a) preparing a sequencing library from a sample comprising a plurality of double-stranded DNA molecules from a patient exposed to the substance, wherein preparing the sequencing library comprises ligating cypher polynucleotides to the double-stranded DNA molecules to form double-stranded cypher-target nucleic acid complexes, wherein the cypher polynucleotides comprise identifier tags selected from a plurality of distinct identifier tag sequences;
- (b) sequencing first and second strands of the cypher-target nucleic acid complexes to produce a plurality of first-strand sequencing reads and a plurality of second-strand sequencing reads;
- (c) for each cypher-target nucleic acid complex among a plurality of the cypher-target nucleic acid complexes, comparing the first-strand sequencing reads with the second-strand sequencing reads to identify nucleotides in the first strand that have a corresponding complementary nucleotide in the second strand;

- (d) detecting a mutation from exposure to the substance by analyzing an error-corrected sequence generated from the first strand sequencing reads and second strand sequencing reads for each of the cypher-target nucleic acid complexes among a plurality of the cypher-target nucleic acid complexes; and comparing the error-corrected sequences to a reference sequence to identify one or more of a mutation, a genomic distribution of mutations, a mutation frequency, sequence heterogeneity, or DNA damage.

40. The method of claim **39**, wherein the sample is derived from a human.

41. The method of claim **39**, wherein prior to preparing the sequencing library, the method further comprises exposing the patient to the substance.

42. The method of claim **41**, wherein prior to exposing the patient to the substance, the patient comprises a normal tissue.

43. The method of claim **39**, wherein the sample comprises a blood sample.

44. The method of claim **39**, wherein the identifier tag sequences comprise random or partially random sequences.